

FIG. 1

FIG. 2A

5' T G G A T G G A G A C C 3'
 80
 FINGER 3 FINGER 2 FINGER 1
FIG. 2B

3/26

$\frac{1}{100}$

$\frac{1}{20}$

2 3

1 2 3 4

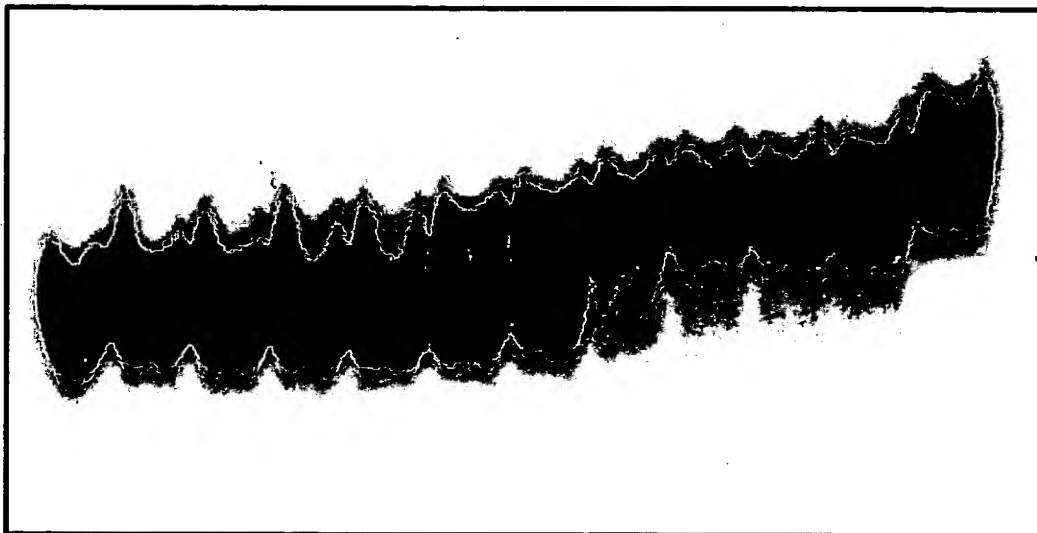


FIG. 3

1:20

0

.5

1.0

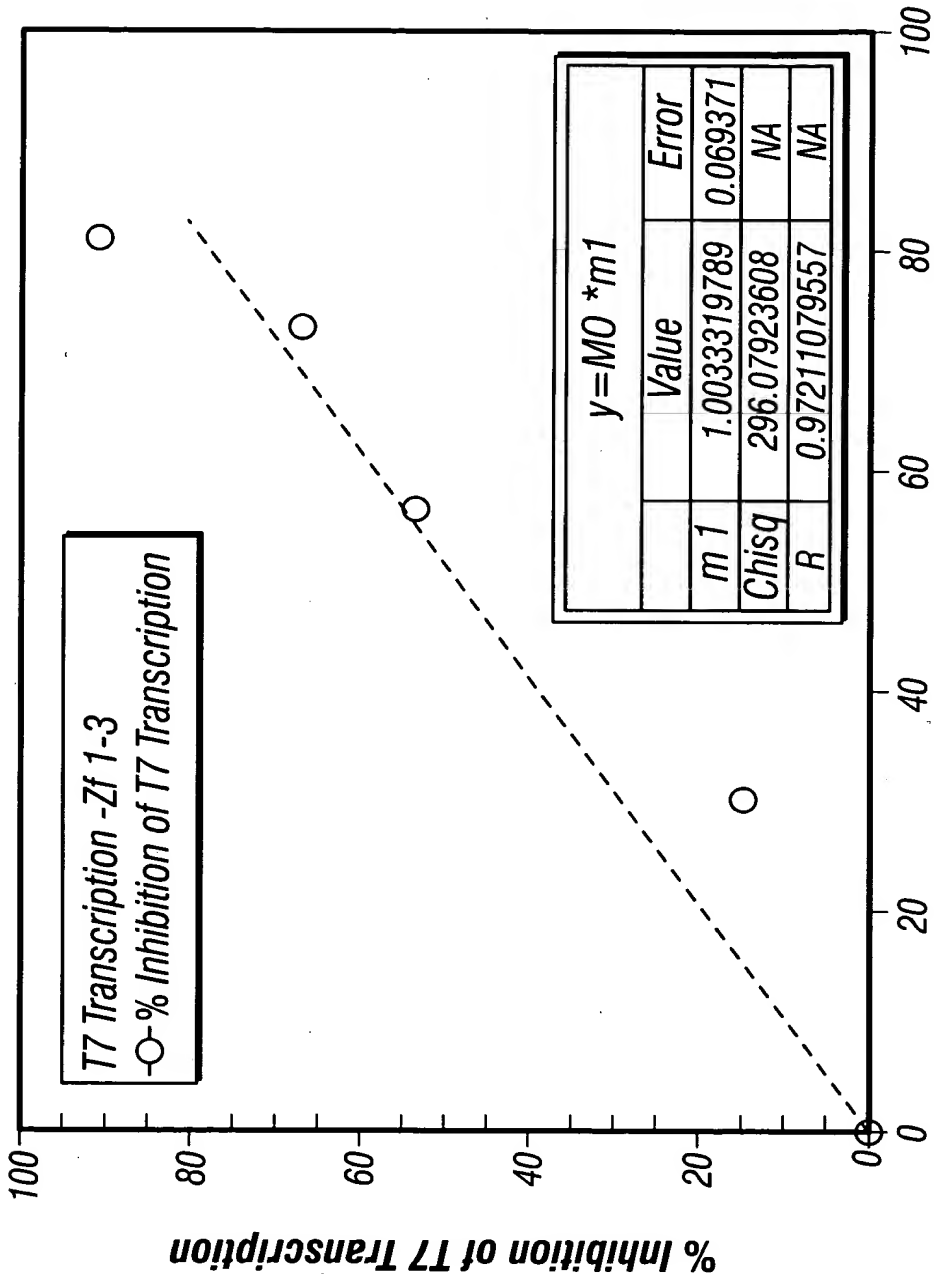
2.0

3.0

4.0



FIG. 4



% DNA Molecules Bound by Zf 1-3

FIG. 5

5/26

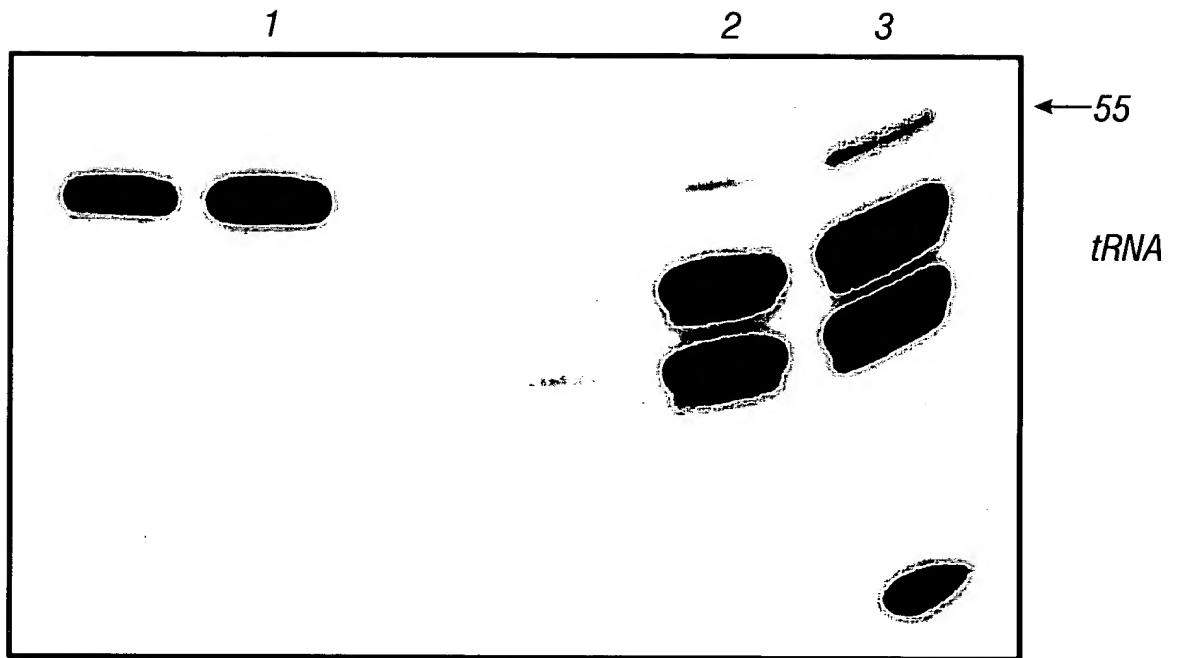


FIG. 6

FIG. 7A

7/26

150	160	170	180	190
CTT ACC ACC CAC ATC CAC ACC CAC ACA GGC GAG AAG CCT TTT GCC TGT				*
GAA TGG TGG GTG TAG GCG TGG GTG TGT CCG CTC TTC GGA AAA CGG ACA				
L T T H I R T E T G E K P F A C>				
200	210	220	230	240
GAC ATT TGT GGT GGC AGG AAG TTT GCC AGG AGT GAT GAA CGC AAG AGG CAT			*	*
CTG TAA ACA CCC TCC TTC AAA CGG TCC TCA CTA CTT GCG TTC TCC GTA				
D I C G R K F A R S D E R K R E>				
250	260	270		
ACG AAA ATC CAT TTA AGA CAG AAG GAC ACT AGT		*		
TGG TTT TAG GTA AAT TCT GTC TTC CTG TGA TCA				
T K I E L R Q K E T S>				

FIG. 7B

8/26

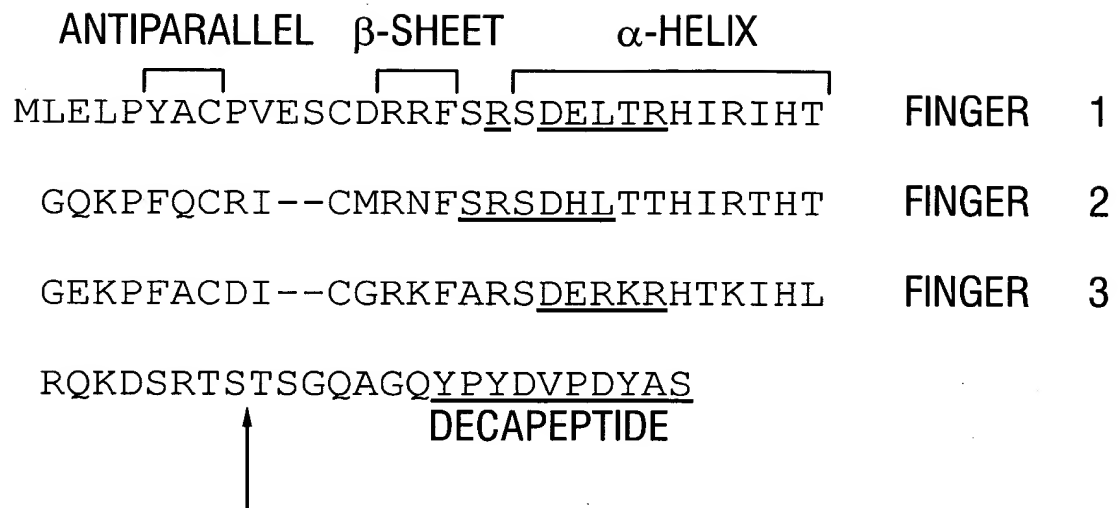


FIG. 8A

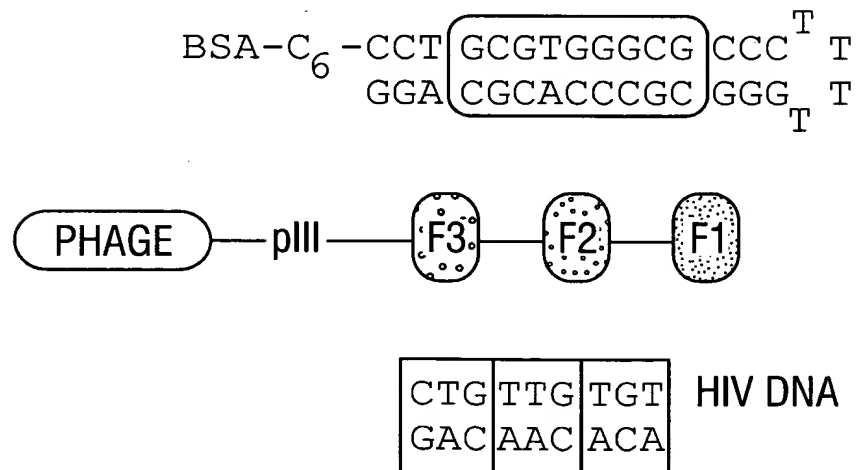


FIG. 8B

9/26

AMINO ACID SEQUENCES OF SELECTED ZINC FINGER PROTEINS								
FINGER	1	SELECTION	FINGER	2	SELECTION	FINGER	3	SELECTION
GCG		TGT	TGG		TTG	GCG		CTG
-123456		-123456	-2-11234		-2-11234	-123456		-123456
RDELTR-(WT)			SRSDDL-(WT)			RDERKR-(WT)		
KADLKR-(C7)		QASKA-(F8)	TYLNTP		GVTMQP-(G3)	RDLANS		NVGDKP
KCVRGR-(C9)		PTHLQT-(F15)	GYRAAP		PQPLSD	SGQWWR-(A14)		SWICGI
KCDRGR		PERTQP	LYRYHL		REQVSR-(G4)	SLLVIA		IAMMEL
KYCRTR		TSEADH	PTLVNA		THMWWMI	VSVRGL		IMMTFF
KQLPWT-(C10)		SEQRYP	VRPHQR		QRMRTL-(G5)			RECRML
KNSQHP		HQQNKP	PFCPYR		QRVGLF			IALLDT
KCQMDS		RGQGMA			LRTGNY-(G6)			NVQGLR
QQVTRT		RARQTG			EREFSL			
TQSQSP		ENSFTD			EKESRG			
VHIQAN		NVMGHD			EGVRKN			
		NRGQRK			TGVNSI			
		SRPSQW			TQARPP			
		TSEADH			THMWWMI			

FIG. 9

10/26

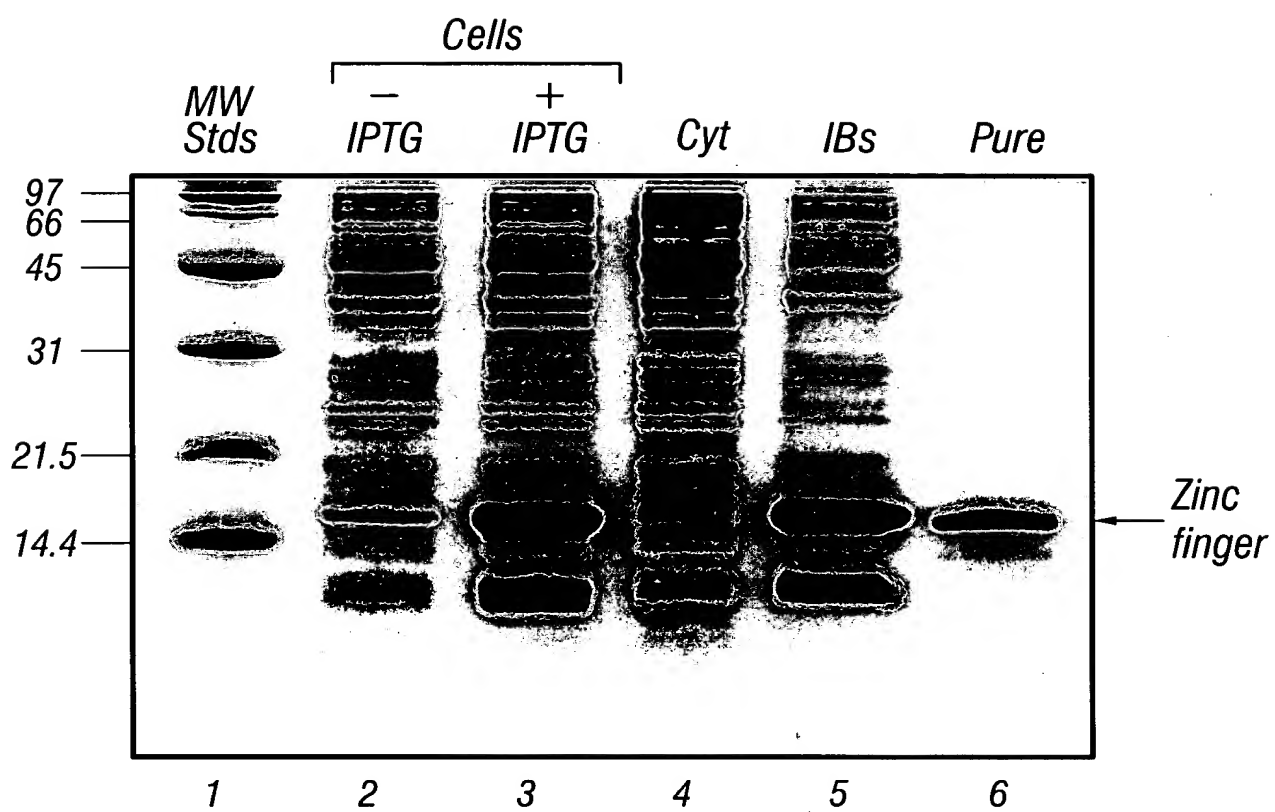


FIG. 10

KINETIC AND EQUILIBRIUM DISSOCIATION CONSTANTS OF ZINC FINGER PROTEINS					
ZINC FINGER	BINDING	$K_{ON}(X10^4)$	$K_{OFF}(X10^{-4})$	$K_d(X10^{-9})$	$K_d/K_d(TARGET)$
PROTEIN	SITE	($M^{-1}s^{-1}$)	(s^{-1})	(M)	
WT	GCG	3.0 ± 0.04	2.0 ± 0.1	6.5	1
	TGT	1.1 ± 0.2	9.0 ± 1.0	81.8	12.6
	GCG	2.4 ± 0.4	1.5 ± 0.7	6.3	
C7	GCG	8.0 ± 0.7	0.4 ± 0.1	0.5	1
	TGT	0.9 ± 0.1	4.9 ± 2.0	54.4	108.8
C9	GCG	2.0 ± 0.2	1.3 ± 0.3	6.5	1
	TGT	0.9 ± 0.1	23.0 ± 3.0	255.6	39.3
C10	GCG	1.8 ± 0.1	4.5 ± 2.0	25.0	1
	TGT	0.3 ± 0.002	1.4 ± 0.1	46.7	1.9
F8	TGT	3.7 ± 1.0	11.0 ± 1.5	29.7	1
	GCG	4.8 ± 0.1	52.0 ± 0.9	108.3	3.6
F15	TGT	1.9 ± 0.1	7.9 ± 1.0	41.6	1
	GCG	0.9 ± 0.3	17.0 ± 1.7	188.9	4.5
G3	TGG	1.7 ± 0.2	2.7 ± 0.2	15.9	1
G4	TGG	2.7 ± 0.3	6.0 ± 0.2	22.2	1.4
	TGG	3.3 ± 0.2	2.1 ± 0.1	6.4	1
	TGG	2.5 ± 0.6	5.7 ± 0.2	22.8	3.6
G5	TGG	0.8 ± 0.1	2.2 ± 0.02	27.5	1
	TGG	1.9 ± 0.2	9.1 ± 0.1	47.9	1.7
G6	TGG	10.0 ± 1.0	4.6 ± 0.3	4.6	1
	TGG	0.7 ± 0.1	1.4 ± 0.1	20.0	4.3
A14	GCG	1.3 ± 0.1	1.7 ± 0.0	13.1	1
	CTG	0.2 ± 0.0	10.0 ± 0.4	500.0	38.2

FIG. 11

12/26

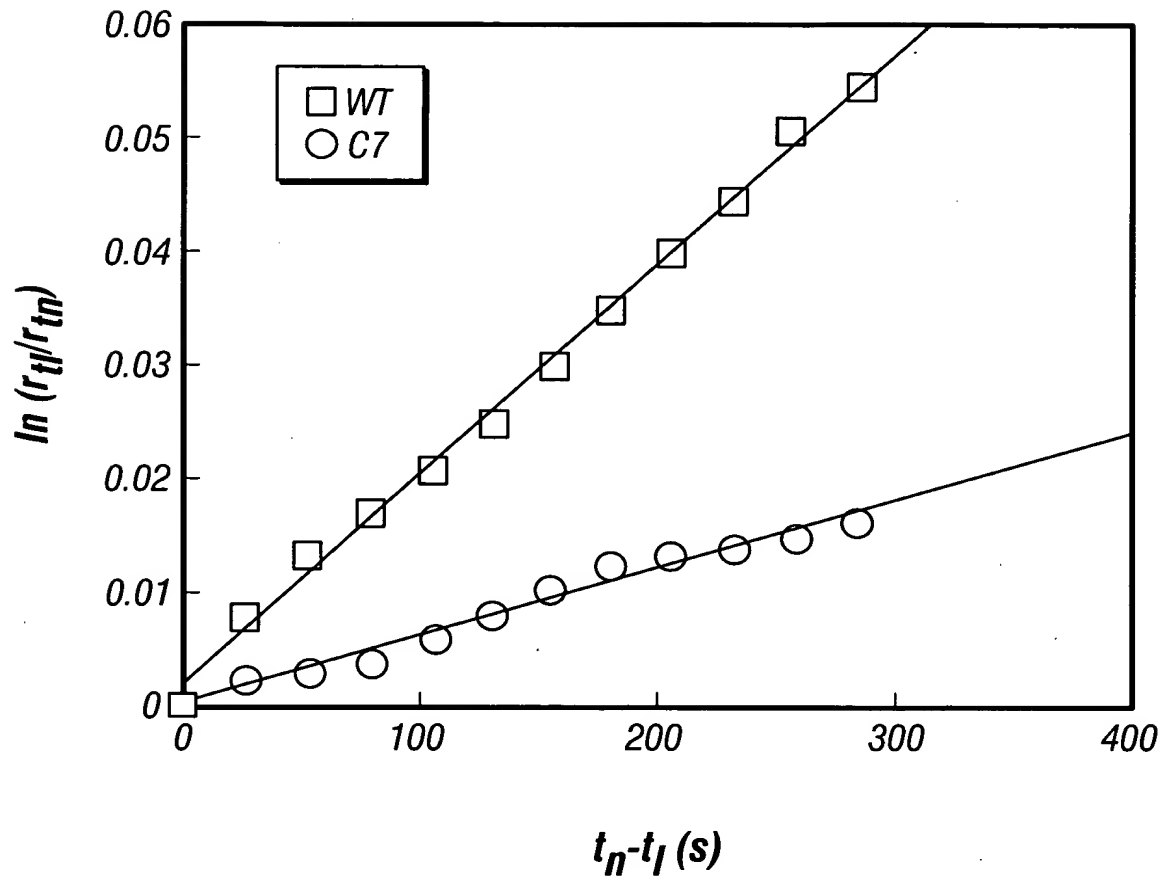


FIG. 12

10	20	30	40	
ATG CTC GAG CTC CCC TAT GCT TGC CCT GTC GAG TCC TGC GAT CGC CGC				
TAC GAG CTC GAG GGG ATA CGA ACG GGA CAG CTC AGG ACG CTA GCG GCG				
M L E L P Y A C C P V E S C D R R>				
50	60	70	80	90
TTT TCT CGC TCG GAT GAG CTT ACC CGC CAT ATC CGC ATC CAC ACA GGC				
AAA AGA GCG AGC CTA CTC GAA TGG GCG GTA TAG GCG TAG GTG TGT CCG				
F S R S D E L T R I R I H T G>				
100	110	120	130	140
CAG AAG CCC TTC CAG TGT CGA ATA TGC ATG CGT AAC TTC AGT CGT AGT				
GTC TTC CCG AAG GTC ACA GCT TAT ACG TAC GCA TTG AAG TCA GCA TCA				
Q K P F Q C R I C M R N F S R S>				
150	160	170	180	190
GAC CAC CTT ACC ACC CAC ATC CGC ACC CAC CAC ACA GGC GAG AAG CCT TTT				
CTG GTG GAA TGG TGG GTG TAG GCG TGG TGG GTG TGT CCG CTC TTC GGA AAA				
D H L T T H I R T T H T G E K P F>				

FIG. 13A-1

14/26

200	210	220	230	240
*	*	*	*	*
GCC TGT GAC ATT TGT GGG AGG AAG TTT GCC AGG AGT GAT GAA CGC AAG				
CGG ACA CTG TAA ACA CCC TCC TTC AAA CGG TCC TCA CTA CTT GCG TTC				
A C D I C G R K F A R S D E R K>				
250	260	270	280	
*	*	*	*	
AGG CAT ACC AAA ATC CAT ACC GGT CAG AAG CCC ACT AGT GGC GGT GGT				
TCC GTA TGG TTT TAG GTA TGG CCA GTC TTC GGG TGA TCA CCG CCA CCA				
R H T K I H T G Q K P T S G G G>				
290	300	310	320	330
*	*	*	*	*
CGG ATC GCC CGG CTG GAG GAG GAA AAA GTG AAA ACC TTG AAA GCG CAA AAC				
GCC TAG CGG GCC GAC CTC CTT TTT CAC TTT TGG AAC TTT CCG GTT TTG				
R I A R R L E E K V K T L K A Q N>				
LINKER				
LJUN				

FIG. 13A-2

340	350	360	370	380
TCC GAG CTG GCG TCC ACC GCC AAC ATG CTC AGG GAA CAG GTG GCA CAG				
AGG CTC GAC CGC AGG TGC CGG TTG TAC GAG TCC CTT GTC CAC CGT GTC				
S E L A S T A N M L R E Q V A Q>				
390	400	410	420	430
CTT AAA CAG AAA GTC ATG AAC CAC GCT AGC GGC CAG GCC GGC CAG TAC				
GAA TTT GTC TTT CAG TAC TTC GTG CGA TCG CCG GTC CGG CCG GTC ATG				
L K Q K V M N H A S G Q A G Q Y>				
440	450	460		
CCG TAC GAC GTT CCG GAC TAC GCT TCT TAA				
GGC ATG CTG CAA GGC CTG ATG CGA AGA ATT				
P Y D V P D Y A S *>				

DECAPEPTIDE TAG

FIG. 13B

10	20	30	40
ATG CTC GAG CTC CCC TAT GCT TGC CCT GTC GAG TCC TGC GAT CGC CGC	*	*	*
TAC GAG CTC GAG GGG ATA CGA ACG GGA CAG CTC AGG ACG CTA GCG GCG			
M L E L P Y A C P V E S C D R R>			
50	60	70	80
TTT TCT CGC TCG GAT GAG CTT ACC CGC CAT ATC CGC ATC CAC ACA GGC	*	*	*
AAA AGA GCG AGC CTA CTC GAA TGG GCG GTA TAG GCG TAG GTG TGT CCG			
F S R S D E L T R H I R I H T G>			
100	110	120	130
CAG AAG CCC TTC CAG TGT CGA ATA TGC ATG CGT AAC TTC AGT CGT AGT	*	*	*
GTC TTC GGG AAG GTC ACA GCT TAT ACG TAC GCA TTG AAG TCA GCA TCA			
Q K P F Q C R I C M R N F S R S>			
150	160	170	180
GAC CAC CTT ACC ACC CAC ATC CGC ACC CAC ACA GGC GAG AAG CCT TTT	*	*	*
CTG GTG GAA TGG TGG GTG TAG GCG TGG GTG TGT CGG CTC TTC GGA AAA			
D H L T T H I R T H T G E K P F>			

FIG. 14A-1

200	210	220	230	240
*	*	*	*	*
GCC TGT GAC ATT TGT GGG AGG AAG TTT GCC AGG AGT GAT GAA CGC AAG				
CGG ACA CTG TAA ACA CCC TCC TTC AAA CGG TCC TCA CTA CTT GCG TTC				
A C D I C G R K F A R S D E R K>				
250	260	270	280	
*	*	*	*	
AGG CAT ACC AAA ATC CAT ACC GGT CAG AAG CCC ACT AGT GGC GGT GGT				
TCC GTA TGG TTT TAG GTA TGG CCA GTC TTC GGG TGA TCA CCG CCA CCA				
R H T K T H T G Q K P T S G G G>				
				LINKER
290	300	310	320	330
*	*	*	*	*
CTG ACC GAC ACC CTG CAG GCG GAA ACC GAC CAG CTG GAA GAC GAA AAA				
GAC TGG CTG TGG GAC GTC CGC CTT TGG CTG GAC CTT CTG CTT TTT				
L T D T L Q A E T D Q Q L E D E K>				

└─ FOS

FIG. 14A-2

440	*		450	*	460	*
CCG TAC GAC GTT CCG GAC TAC GCT TCT TAA						
GGC ATG CTG CAA GGC CTG ATG CGA AGA ATT						
P Y D V P D Y A S >						

DECAPEPTIDE TAG

FIG. 14B

10	20	30	40
*	*	*	*
ATG AAA CTG CTC GAG CCC TAT GCT TGC CCT GTC GAG TCC TGC GAT CGC			
TAC TTT GAC GAG CTC GGG ATA CGA ACG GGA CAG CTC AGG ACG CTA GCG			
M K L L E P Y A C P V E S C D R>			
50	60	70	80
*	*	*	*
CGC TTT TCT AAG TCG GCT GAT CTG AAG CGC CAT ATC CGC ATC CAC ACT			
GCG AAA AGA TTC AGC CGA CTA GAC TTC GCG GTA TAG GCG TAG GTG TGA			
R F S K S A D L K R H I R I H T>			
100	110	120	130
*	*	*	*
GGC GAA AAA CCG TAC GCG TGC CCT GTC GAG TCC TGC GAT CGC CGC TTT			
CCG CTT TTT GGC ATG CGC ACG GGA CAG CTC AGG ACG CTA GCG GCG AAA			
G E K P Y A C P V E S C D R R F>			
150	160	170	180
*	*	*	*
TCT AAG TCG GCT GAT CTG AAG CGC CAT ATC CGC ATC CAC ACC GGG GAG			
AGA TTC AGC CGA CTA GAC TTC GCG GTA TAG GCG TAG GTG TGG CCC CTC			
S K S A D L K R H I R I H T G E>			

FIG. 15A

250 * 260 * 270 * 280 *

TCG GCT GAT CTG AAG CGC CAT ATC CGC ATC CAC ACC GGT CAG AAG CCC
AGC CGA CTA GAC TTC GCG GTA TAG GCG TAG GTG TGG CCA GTC TTC GGG
S A D L K R R H I R I N T G Q K P>

290

FIG. 15B

FIG. 16A-1

FIG. 16A-2

[illegible]

	390	400	410	420	430
CGT AAC	TTC AGT	CGT AGT	GAC CAC	CTT ACC	ACC CAC
GCA TTG	AAG TCA	GCA TCA	CTG CTG	GAA TGG	TGG TAC
R N	F S	R S	D H	L T	I R
					T H
					T H

	440		450		460		470		480						
	*		*		*		*		*						
ACA	GGC	GAG	AAG	CCT	TTT	GCC	TGT	GAC	ATT	TGT	GGG	AGG	AAG	TTT	GCC
TGT	CCG	CTC	TTC	GGA	AAA	CGG	ACA	CTG	TAA	ACA	CCC	TCC	TTC	AAA	CGG
T	G	E	K	P	F	A	C	D	I	C	G	R	K	F	A

	490	500	510	520
	*	*	*	*
AGG AGT GAT GAA CGC AAG AGG CAT ACC AAA ATC CAT TTA AGA CAG AAG				
TCC TCA CTA CTT GCG TTC TCC GTA TGG TTT TAG GTA AAT TCT GTC TTC				
R S D E R K R R H T K I H L R O K>				

530	*	GAC	TCT	AGA	ACT	AGT
		CTG	AGA	TCT	TGA	TCA
		D	S	R	T	S>

FIG. 16B

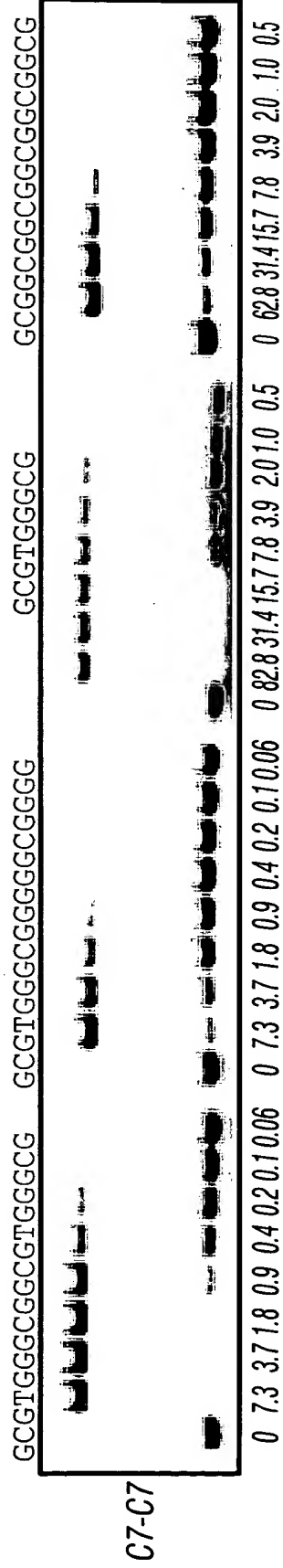
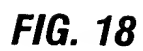


FIG. 17A



FIG. 17B



26/26

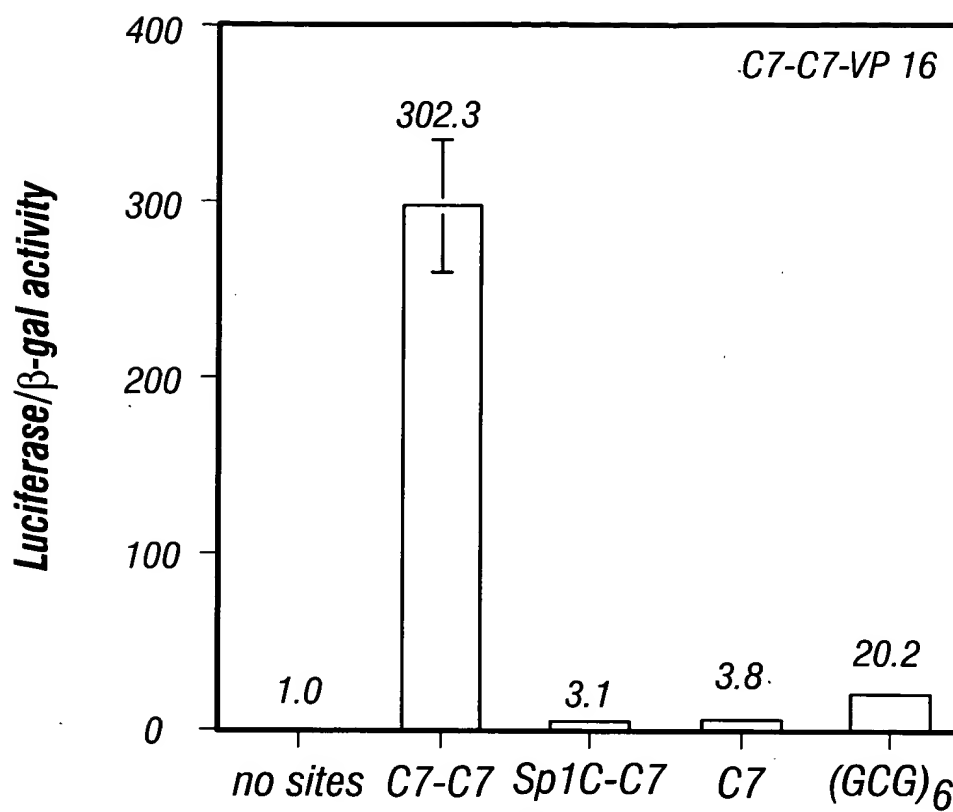


FIG. 19A

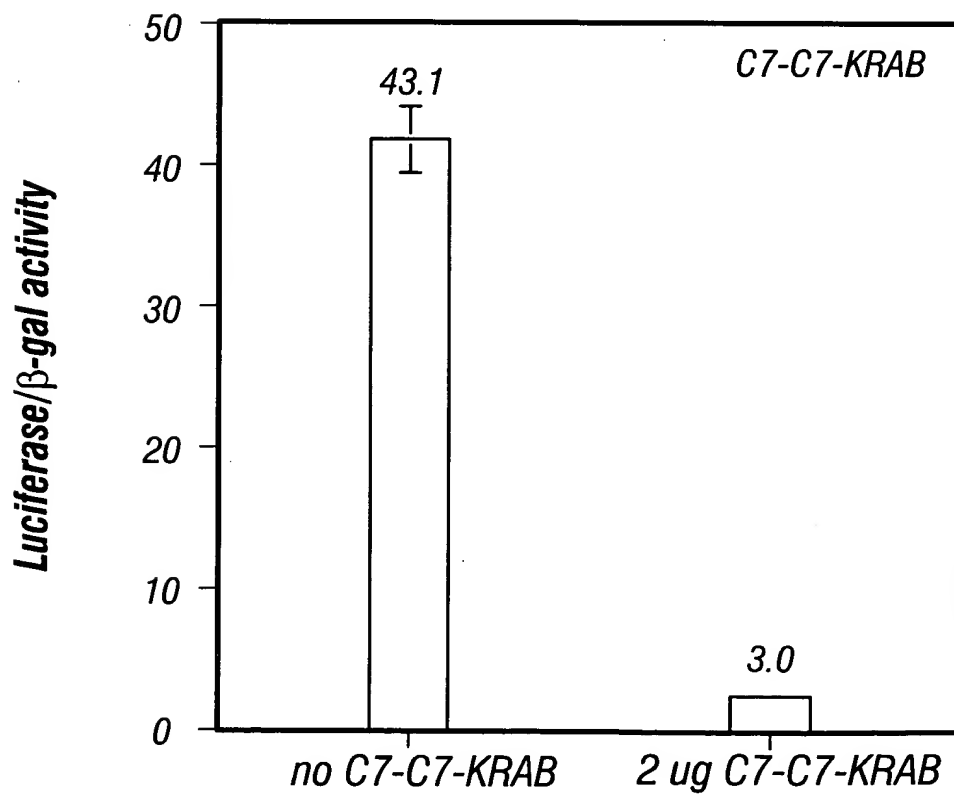


FIG. 19B